SEQUENCE LISTING

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<110> Fainzilber, Michael M.
             Kits, Karel S.
             Burlingame, Alma L.
             Olivera, Baldomero M.
             Walker, Craig
             Watkins, Maren
             Shetty, Reshma
             Cruz, Lourdes J.
DEC 0 6 1999
             Imperial, Julita
             Colledge, Clark
             University of Utah Resarch Foundation
             Vrije Universiteit
             Regents of the University of California
       <120> Gamma-Conopeptides
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       <151> 1997-12-16
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       <223> Xaa at residues 1, 2, 3, 4, 5, and 6 may be
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Xaa Xaa Xaa Cys Cys Xaa Xaa Xaa Xaa Cys Xaa Xaa Xaa Xaa Xaa
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      and 39 may be des-Xaa or any amino acid.
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    _13 may be any amino acid.
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Xaa Xaa

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       or hydroxy-Pro.
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 Ser Asn Ser Cys Asp Gln Thr Tyr Cys Xaa Leu Tyr Ala Phe Xaa Ser
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 <213> Conus textile
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 <221> PEPTIDE
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       are Pro or hydroxy-Pro.
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Cys Cys Ser Asn Ser Cys Asp Gln Thr Tyr Cys Thr Leu Met Xaa Xaa
 Asp Xaa
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       37 ar Pro or hydroxy-Pro.
```

Xaa Xaa Arg Xaa Gly Gly Cys Met Ala Xaa Phe Gly Leu Cys Ser Arg

36

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1
                                        10
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Met Xaa Phe Xaa Xaa Asp Xaa
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      or gamma-carboxyglutamate; Xaa at residue 12 is
      Pro or hydroxy-Pro.
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Xaa Val
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Ser Xaa Cys Cys Ser Xaa Asp Cys Xaa Gly Ser Cys Thr Met Xaa
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Ser Leu Leu Met Cys Ser Lys Ala Thr Ser Arg Cys Ile Leu Ala Leu
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Arg Trp Leu Glu Cys Ser Val Trp Phe Ser His Cys Thr Lys Asp Ser
             20
gaa tgt tgt tct aat agt tgt gac caa acg tac tgc acg tta atg cca
Glu Cys Cys Ser Asn Ser Cys Asp Gln Thr Tyr Cys Thr Leu Met Pro
         35
ccg gac tgg tgacatcgcc actctcctgt tcagagtctt caaggctttt
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Pro Asp Trp
    50
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Leu	Ser	Lys 35	Arg	Lys	Pro	Pro	Ala 40	Glu	Arg	Trp	Trp	Arg 45	Trp	Gly	Gly	
Cys	Met 50	Ala	Trp	Phe	Gly	Leu 55	Cys	Ser	Arg	Asp	Ser 60	Glu	Cys	Cys	Ser	
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Leu Thr Ile Leu Leu Val Ala Ala Val Leu Met Ser Thr Gln Ala
cta att caa gat caa cgc caa aag gca aag atc aac ttg ttt tca aag
                                                                   153
Leu Ile Gln Asp Gln Arg Gln Lys Ala Lys Ile Asn Leu Phe Ser Lys
20
                                         30
aga cag gca tat gct cgt gat tgg tgg gac gat ggc tgc agt gtg tgg
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Arg Gln Ala Tyr Ala Arg Asp Trp Trp Asp Asp Gly Cys Ser Val Trp
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ggg cct tgt acg gtg aac gca gaa tgt tgt tct ggt gat tgt cat gaa
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Gly Pro Cys Thr Val Asn Ala Glu Cys Cys Ser Gly Asp Cys His Glu
acg tgc att ttc ggg tgg gaa gtc tgaccacaaa ccatccgaca tcgccactct
Thr Cys. Ile Phe Gly Trp Glu Val
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Phe	Ser	Lys 35	Arg	Gln	Ala	Tyr	Ala 40	Arg	Asp	Trp	Trp	Asp 45	Asp	Gly	Cys	
Ser	Val 50	Trp	Gly	Pro	Cys	Thr 55	Val	Asn	Ala	Glu	Cys 60	Cys	Ser	Gly	Asp	
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			cgt Arg													214
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Leu Ile Gln Gly Gly Gly Asp Lys Arg Gln Lys Ala Asn Ile Asn Phe
ctt tca agg tgg gac cgt gag tgc agg gct tgg tat gcg ccg tgt agc
Leu Ser Arg Trp Asp Arg Glu Cys Arg Ala Trp Tyr Ala Pro Cys Ser
cct ggc gcg caa tgt tgt agt ttg ctg atg tgt tca aaa gcg acc agc
Pro Gly Ala Gln Cys Cys Ser Leu Leu Met Cys Ser Lys Ala Thr Ser
                         55
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cgc tgc ata ttg gcg tta tgaactctga ccacaagcca tccgacatca
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Arg Cys Ile Leu Ala Leu
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tgaacaaata agtagaatag cacgtttttc cccctttgca aaatcaataa tggaggttaa 361
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44

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Leu	Ser	Arg 35	Trp	Asp	Arg	Glu	Cys 40	Arg	Ala	Trp	Tyr	Ala 45	Pro	Cys	Ser	
Pro	Gly 50	Ala	Gln	Cys	Cys	Ser 55	Leu	Leu	Met	Cys	Ser 60	Lys	Ala	Thr	Ser	
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			aaa Lys 35													144
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			atg Met							tgaa	actct	iga (ccaca	aagco	ca	242
tcc	gacat	ca d	ccact	ctc	et et	tcaç	gagto	tto	caag	(8 +	An i americo			- +	278
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			Leu	Ile 5	lle	Leu	Leu	Leu	Val 10	Ala	Ala	Val	Leu	Leu 15	Ser	
Thr	Gln	Ala	Leu 20	Asn	Gln	Glu	Lys	Arg 25	Pro	Lys	Glu	Met	Ile 30	Asn	Phe	,
Leu	Ser	Lys	Gly	Lys	Thr	Asn	Ala	Glu	Arg	Arg	Asn	Gly	Gln	Cys	Glu	

Asp Val Trp Met Pro Cys Thr Ser Asn Trp Glu Cys Cys Ser Leu Asp 50 55.

Cys Glu Met Tyr Cys Thr Gln Ile Gly

<220>

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<211> 287
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<220>
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    Met Glu Lys Leu Thr Ile Leu Leu Leu Val Ala Val Leu Ile
eeg ace eag gee ett ttt eaa ggt gat gae gga aaa tee eag aag geg
                                                                   96
Pro Thr Gln Ala Leu Phe Gln Gly Asp Asp Gly Lys Ser Gln Lys Ala
gag atc aag tct ttt gaa aca aga aag tta gcg aga aac aag cag gta
                                                                   144
Glu Ile Lys Ser Phe Glu Thr Arg Lys Leu Ala Arg Asn Lys Gln Val
cgc tgc ggt ggt tgg tca acg tat tgt gaa gtt gac gag gaa tgc tgt
                                                                   192
Arg Cys Gly Gly Trp Ser Thr Tyr Cys Glu Val Asp Glu Glu Cys Cys
         50
tcg gaa tca tgt gta agg tct tac tgc acg ctg ttt gga tgaactcgga
                                                                   241
Ser Glu Ser Cys Val Arg Ser Tyr Cys Thr Leu Phe Gly
ccacaagcca tccgatatca ccactctcct gttcagagtc ttcaag
                                                                   287
<210> 31
<211> 76
<212> PRT
<213> Conus marmoreus
<400> 31
Met Glu Lys Leu Thr The Leu Leu Val Ala Ala Val Leu Ile Pro
Thr Gln Ala Leu Phe Gln Gly Asp Asp Gly Lys Ser Gln Lys Ala Glu
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Ile Lys Ser Phe Glu Thr Arg Lys Leu Ala Arg Asn Lys Gln Val Arg
Cys Gly Gly Trp Ser Thr Tyr Cys Glu Val Asp Glu Glu Cys Cys Ser
Glu Ser Cys Val Arg Ser Tyr Cys Thr Leu Phe Gly
<210> 32
<211> 278
<212> DNA
<213> Conus marmoreus
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46

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<221> CDS
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<400> 32
atc atg cag aaa ctg ata att ctg ctt ctt gtt gct gct gtg ctg atg
                                                                   48
   Met Gln Lys Leu Ile Ile Leu Leu Val Ala Ala Val Leu Met
acg acc cag gcc cta tat caa gaa aaa cgc cga aag gag atg atc aat
                                                                   96
Thr Thr Gln Ala Leu Tyr Gln Glu Lys Arg Arg Lys Glu Met Ile Asn
                 20
                                     25
ttt tta tca aaa gga aag ata aat gct gag agg cgg aac ggc gga tgc
                                                                   144
Phe Leu Ser Lys Gly Lys Ile Asn Ala Glu Arg Arg Asn Gly Gly Cys
             35
                                 40
aaa get act tgg atg tet tgt tea teg gge tgg gaa tge tgt tet atg
                                                                   192
Lys Ala Thr Trp Met Ser Cys Ser Ser Gly Trp Glu Cys Cys Ser Met
         50
                             55
agt tgt gac atg tac tgc gga tagataggat gaactctgac cacaagccat
                                                                   243
Ser Cys Asp Met Tyr Cys Gly
ccgacatcac cactctcctc ttcagagtct tcaag
                                                                   278
<210> 33
<211> 70
<212> PRT
<213> Conus marmoreus
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Met Gln Lys Leu Ile Ile Leu Leu Val Ala Ala Val Leu Met Thr
Thr Gln Ala Leu Tyr Gln Glu Lys Arg Arg Lys Glu Met Ile Asn Phe
Leu Ser Lys Gly Lys Ile Asn Ala Glu Arg Arg Asn Gly Gly Cys Lys
Ala Thr Trp Met Ser Cys Ser Ser Gly Trp Glu Cys Cys Ser Met Ser
Cys Asp Met Tyr Cys Gly
65
<210> 34
<211> 528
<212> DNA
<213> Conus textile
<220>
<221> CDS
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geacgteate ttetetetea gtetgeetga eagetgeett eagteaacce tgeegteate 60
tcagogtaga ottggtaaga agtgaaaaao atttato_atg cag aaa otg ata ato_ 115
                                         Met Gln Lys Leu Ile Ile
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ctg ctt ctt gtt g Leu Leu Leu Val A 10				
gaa aaa cgc cca a Glu Lys Arg Pro I 25		Lys Leu Leu		
gat gct gag aag c Asp Ala Glu Lys G 40				
tgt tca cat gcc c Cys Ser His Ala H 55				
tgt acg gga tgaac Cys Thr Gly	tcgga ccacaagc	ca teegaeatea	a ccactctcct	356
cttcagaggc ttcaag	actt ttgttctga	t tttggacaat	ctttacgagt aaac	caaataa 416
ttagactagc actttt	tttc ccctttgca	a aatcaatgat	ggaggtaaaa agco	ctcccat 476
tttgtcttca tcaata	aaga acttatcat	c aaaaaaaaa	aaaaaaaaa aa	528
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Thr Gln Ala Val L 20	eu Gln Glu Lys	Arg Pro Lys 25	Glu Lys Ile Lys 30	s Leu
Leu Ser Lys Arg L 35	ys Thr Asp Ala 40	-	Gln Lys Arg Let 45	ı Cys
Pro Asp Tyr Thr G	lu Pro Cys Ser	His Ala His	Glu Cys Cys Ser 60	Trp
Asn Cys Tyr Asn G 65	ly His Cys Thr 70	Gly		
<210> 36 <211> 26 <212> PRT <213> Conus texti	le			
gamma-carbo	due 18 is Trp and 14 are Glu xyglutamate; X hydroxy-Pro.	or	-	
<400> 36 Leu Cys Xaa Asp T 1	yr Thr Xaa Xaa 5	Cys Ser His	Ala His Xaa Cys	

Ser Xaa Asn Cys Tyr Asn Gly His Cys Thr

20

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<210> 37
<211> 4
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<213> Artificial Sequence
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<223> Description of Artificial Sequence:consensus
      gamma-conopeptide sequence for probe
<221> PEPTIDE
<222> (1)
<223> Xaa is Glu or Gln.
<400> 37
Xaa Cys Cys Ser
  1.
<210> 38
<211> 12
<212> DNA
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<223> Description of Artificial Sequence:degenerate
      probe for consensus gamma-conopeptide sequence.
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                                                                    12
sartgytgya gy
<210> 39
<211> 12
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: degenerate
      probe for consensus gamma-conopeptide sequence.
<400> 39
                                                                    12
sartgytgyt cn
<210> 40
<211> 8
<212> PRT
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<223> Description of Artificial Sequence:consensus
      pro-gamma-conopeptide sequence for probe.
<400> 40
Ile Leu Leu Val Ala Ala Val Leu
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<210> 41
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<211> 24
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:degenerate
      probe for consensus pro-gamma-conopeptide
      sequence.
<400> 41
                                                                    24
athytnytng tngcngcngt nytn
<210> 42
<211> 32
<212> PRT
<213> Conus pennaceus
<220>
<221> PEPTIDE
<222> (1)..(31)
<223> Xaa at residues 14 and 26 are
      gamma-carboxyglutamate; Xaa at residue 31 is
      hdroxy-Pro.
<400> 42
Asp Cys Thr Ser Trp Phe Gly Arg Cys Thr Val Asn Ser Xaa Cys Cys
Ser Asn Ser Cys Asp Gln Thr Tyr Cys Xaa Leu Tyr Ala Phe Xaa Ser
             20
<210> 43
<211> 27
<212> PRT
<213> Conus textile
<220>
<221> PEPTIDE
<222> (1)..(27)
<223> Xaa at residues 9 and 13 are
      gamma-carboxyglutamate.
<400> 43
Cys Gly Gly Tyr Ser Thr Tyr Cys Xaa Val Asp Ser Xaa Cys Cys Ser
Asp Asn Cys Val Arg Ser Tyr Cys Thr Leu Phe
             20
                                  25
<210> 44
<211> 8
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<213> Conus pennaceus
<220>
<221>_ MOD RES
<222> (2)
<223> Xaa at residue 2 is carboxymethylCys
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<400> 44
Asp Xaa Thr Ser Trp Phe Gly Arg
<210> 45
<211> 24
<212> PRT
<213> Conus pennaceus
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<221> PEPTIDE
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<223> Xaa at residues 6 and 18 are
      gamma-carboxyglutamate; Xaa at residue 23 is
<400> 45
Xaa Thr Val Asn Ser Xaa Xaa Xaa Ser Asn Ser Xaa Asp Gln Thr Tyr
                                      10
Xaa Xaa Leu Tyr Ala Phe Xaa Ser 20
<210> 46
<211> 18
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: primer for M13
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<400> 46
                                                                    18
tttcccagtc acgacgtt
<210> 47
<211> 19
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence:primer for M13
      reverse priming site.
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<400> 47

cacacaggaa acagctatg

51

19